

## INFORMATION DISCLOSURE STATEMENT

(Use several sheets if necessary)

ATTY. DOCKET NO.  
690068.405C3APPLICATION NO.  
10/015,540APPLICANTS  
Nicholas Ling et al.FILING DATE  
December 11, 2001GROUP ART UNIT  
Not Yet Assigned

## U.S. PATENT DOCUMENTS

*EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
ms	AA	5,288,639	2/22/94	Burnie et al.	435	320.1	
	AB	5,468,481	11/21/95	Sharma et al.	530	402	
	AC	5,559,209	9/24/96	Nishimoto	530	326	
	AD	5,948,764	9/7/99	Gaur et al.	514	14	
ms	AE	6,036,957	3/14/00	Weiner et al.	424	184.1	

## FOREIGN PATENT DOCUMENTS

		DOCUMENT NUMBER	DATE	COUNTRY	TRANSLATION	
					YES	NO
ms	AF	JP 6157591	6/3/94	Japan		
	AG	EP 304 279 B1	2/22/89	EPO		
	AH	WO 88/00057	1/14/88	WIPO		
	AI	WO 88/10120	12/29/88	WIPO		
ms	AJ	WO 91/12816	9/5/91	WIPO		

## OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

ms	AK	Abstract 65027 from <i>Biol. Abstr.</i> 81(7): AB-701 (1985) of the article "Monoclonal antibodies to human myelin basic protein," <i>J. Neurochem.</i> 46(1): 47-53, 1985.
	AL	Acha-Orbea et al., "Limited Heterogeneity of T Cell Receptors from Lymphocytes Mediating Autoimmune Encephalomyelitis Allows Specific Immune Intervention," <i>Cell</i> 54: 263-273, 1988.
	AM	Antel et al., "Vaccination with an altered MBP Peptide Induces a Persistent Immune Response," <i>Journal of Neuroimmunology</i> 90(1): p. 18, Abstract No. 80, 1998.
	AN	Arimilli et al., "Identification of Core Structure and Critical T Cell Receptor Contact Residues in an Antigenic Peptide by Measuring Acidification Rates," <i>Journal of Immunological Methods</i> 217: 49-59, 1998.
ms	AO	Ausubel et al., "Changes in Cytokine Secretion Induced by Altered Peptide Ligands of Myelin Basic Protein Peptide 85-99," <i>Journal of Immunology</i> 159: 2502-2512, 1997.

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FORM PTO-1449 (REV. 7-80)	U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE	ATTY. DOCKET NO. 690068.405C3	APPLICATION NO. 10/015,540	COPY OF PAPERS ORIGINALLY FILED
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					YES	NO
MW	BA	WO 92/06708	4/30/92	WIPO		
	BB	WO 92/21367	12/10/92	WIPO		
	BC	WO 93/08212	4/29/93	WIPO		
	BD	WO 93/12814	7/8/93	WIPO		
MW	BE	WO 93/19178	9/30/93	WIPO		

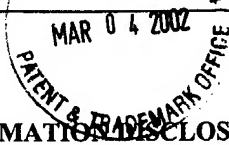
## OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

MW	BF	Babbitt et al., "Antigenic competition at the level of peptide-Ia-binding," <i>Proc. Natl. Acad. Sci. USA</i> 83: 4509-4513, 1986.
	BG	Bellacosa et al., "A Retroviral Oncogen, <i>akt</i> , Encoding a Serine-Threonine Kinase Containing an SH2-Like Region," <i>Science</i> 254(5029): 274-277, 1991.
	BH	Bernard, "Experimental Autoimmune Encephalomyelitis in Mice: Genetic Control of Susceptibility," <i>Journal of Immunogenetics</i> 3: 263-274, 1976.
	BI	Brocke et al., "In Vitro Proliferative Responses and Antibody Titers Specific to Human Acetylcholine Receptor Synthetic Peptides in Patients with Myasthenia Gravis and Relation to HLA Class II Genes," <i>J. Clin. Invest.</i> 82: 1894-1900, 1988.
	BJ	Brocke et al., "Treatment of experimental encephalomyelitis with a peptide analogue of myelin basic protein," <i>Nature</i> 379: 343-346, 346, 1996.
	BK	Brostoff and Howell, "T Cell Receptors, Immunoregulation, and Autoimmunity," <i>Clinical Immunology and Immunopathology</i> 62(1): 1-7, 1992.
	BL	Carter and Rodriguez, "Immunosuppressive Treatment of Multiple Sclerosis," <i>Mayo Clin. Proc.</i> 64: 664-669, 1989.
	BM	Chou et al., "Identity of Myelin Basic Protein from Multiple Sclerosis and Human Control Brains: Discovery of a Genetic Variant," <i>Journal of Neurochemistry</i> 30: 745-750, 1978.
MW	BN	Day et al., "The Polyclonal Antibody Responses of Lewis Rats to the Synthetic Encephalitogenic Neuropeptide S55S (Residues 72-84 of Guinea Pig Myelin Basic Protein) and Its Analogs," <i>Journal of Neuroscience Research</i> 18: 214-221, 1987.

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MB	CA	WO 93/21222	10/28/93	WIPO		
	CB	WO 94/06828	3/31/94	WIPO		
	CC	WO 95/08572	3/30/95	WIPO		
	CD	WO 96/12731	5/2/96	WIPO		
MM	CE	WO 96/16085	5/30/96	WIPO		

## OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

MM	CF	Einstein et al., "Suppression of Experimental Allergic Encephalomyelitis By Chemically Modified Encephalitogen," <i>Immunochemistry</i> 9: 1013-1019, 1972.
	CG	Evavold and Allen, "Separation of IL-4 Production from Th Cell Proliferation by an Altered T Cell Receptor Ligand," <i>Science</i> 252: 1308-1310, 1991.
	CH	Gammon et al., "Neonatal T-cell tolerance to minimal immunogenic peptides is caused by clonal inactivation," <i>Nature</i> 319: 413-415, 1986.
	CI	Gaur et al., "Amelioration of relapsing experimental autoimmune encephalomyelitis with altered myelin basic protein peptides involves different cellular mechanisms," <i>Journal of Neuroimmunology</i> 74: 149-158, 1997.
	CJ	Gaur et al., "Amelioration of Autoimmune Encephalomyelitis by a Myelin Basic Protein Synthetic Peptide-Induced Anergy," <i>Science</i> 258(5087): 1491-1494, 1992.
	CK	Gautam et al., "A Polyalanine Peptide With only Five Native Myelin Basic Protein Residues Induces Autoimmune Encephalomyelitis," <i>J. Exp. Med.</i> 176: 605-609, 1992.
	CL	Gautam et al., "Inhibition Of Experimental Autoimmune Encephalomyelitis By A Nonimmunogenic Non-Self Peptide That Binds To I-A <sup>u</sup> ," <i>The Journal Of Immunology</i> 148(10): 3049-3054, 1992.
MM	CM	Hadden et al., "Thymic Hormones, Interleukins, Endotoxin and Thymomimetic Drugs in T Lymphocyte Ontogeny," in <i>Advances in Immunopharmacology</i> 3, L. Chedid et al. (Eds.), 1985, pp. 487-497.

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					YES	NO
mm	DA	WO 96/16086	5/30/96	WIPO		
	DB	WO 96/28470	9/19/96	WIPO		
	DC	WO 96/34622	11/7/96	WIPO		
mm	DD	WO 97/35879	10/2/97	WIPO		

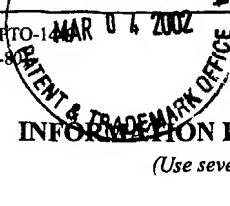
## OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

mm	DE	Hashim, "Experimental Allergic Encephalomyelitis: Activation of Suppressor T Lymphocytes by a Modified Sequence of the T Effector Determinant," <i>Journal of Immunology</i> 126(2): 419-423, 1981.
	DF	Hashim and Day, "Synthetic Peptide Analogs to Probe the Immunological Expression of the Rat Encephalitogenic Neuropeptide," <i>Journal of Neuroscience Research</i> 18: 209-213, 1987.
	DG	Hashim et al., "Suppression and Reversal of Allergic Encephalomyelitis in Guinea Pigs with a Non-Encephalitogenic Analogue of the Tryptophan Region of the Myelin Basic Protein," <i>Journal of Immunology</i> 116(1): 126-130, 1976.
	DH	Jahnke et al., "Sequence Homology Between Certain Viral Proteins and Proteins Related to Encephalomyelitis and Neuritis," <i>Science</i> 229: 282-284, 1985.
	DI	Kardys and Hashim, "Experimental Allergic Encephalomyelitis in Lewis Rats: Immunoregulation of Disease By a Single Amino Acid Substitution in the Disease-Inducing Determinant," <i>Journal of Immunology</i> 127(3): 862-866, 1981.
	DJ	Karin et al., "Reversal of Experimental Autoimmune Encephalomyelitis by a Soluble Peptide Variant of a Myelin Basic Protein Epitope: T Cell Receptor Antagonism and Reduction of Interferon $\gamma$ and Tumor Necrosis Factor $\alpha$ Production," <i>J. Exp. Med.</i> 180: 2227-2237, 1994.
	DK	Kira et al., "Experimental Allergic Encephalomyelitis in Rabbits. A Major Encephalitogenic Determinant within Residues 1-44 of Myelin Basic Protein," <i>J. of Neuroimmunol.</i> 12(3): 183-193, 1986.
mm	DL	Kuchroo et al., "A Single TCR Antagonist Peptide Inhibits Experimental Allergic Encephalomyelitis Mediated by a Diverse T Cell Repertoire," <i>Journal of Immunology</i> 153: 3326-3336, 1994.

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FORM PTO-101 (REV. 7-8)		U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE		ATTY. DOCKET NO. 690068.405C3		APPLICATION NO. 10/015,540	
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<b>OTHER PRIOR ART</b> (Including Author, Title, Date, Pertinent Pages, Etc.)							
MMS	EA	Lamont et al., "Inhibition of Experimental Autoimmune Encephalomyelitis Induction In SJL/J Mice Using A Peptide With High Affinity For IA <sup>S</sup> Molecules," <i>Journal of Immunology</i> 145(6): 1687-1693, 1990.					
	EB	Lehninger, "The Amino Acid Building Blocks of Proteins," in <i>Biochemistry</i> , 2 <sup>nd</sup> ed., Worth Publishers, Inc. pp. 71-75, 1975.					
	EC	Lindsey et al., "Double-Blind, Randomized, Placebo-Controlled Evaluation of the Safety, Tolerability, and Pharmacokinetics of CGP 77116 in Patients with Multiple Sclerosis," <i>Neurology</i> 50: Abstract 149, 1998.					
	ED	Martin et al., "A Myelin Basic Protein Peptide is Recognized by Cytotoxic T Cells in the Context of Four HLA-DR Types Associated with Multiple Sclerosis," <i>Journal of Experimental Medicine</i> 173: 19-24, 1991.					
	EE	Martin et al., "Diversity in Fine Specificity and T Cell Receptor Usage of the Human CD4+ Cytotoxic T Cell Response Specific for the Immunodominant Myelin Basic Protein Peptide 87-106," <i>Journal of Immunology</i> 148(5): 1359-1366, 1992.					
	EF	Martin et al., "Immunological Aspects of Demyelinating Diseases," <i>Annu. Rev. Immunol.</i> 10: 153-187, 1992.					
	EG	Pautot et al., "Leucine aminopeptidase: An inducible component of the defense response in <i>Lycopersicon esculentum</i> (tomato)," <i>Proc. Natl. Acad. Sci. USA</i> 90: 9906-9910, 1993.					
	EH	Raine and Stone, "Animal Model for Multiple Sclerosis. Chronic experimental allergic encephalomyelitis in inbred guinea pigs," <i>New York State Journal of Medicine/September</i> : 1693-1697, 1977.					
	EI	Rothbard, "Peptides and the Cellular Immune Response," <i>Ann. Inst. Pasteur/Virologie</i> 137 E: 518-526, 1986.					
	EJ	Rothbard and Taylor, "A sequence pattern common to T cell epitopes," <i>EMBO J.</i> 7(1): 93-100, 1988.					
MMS	EK	Rudinger, "Characteristics of the amino acids as components of a peptide hormone sequence," in <i>Peptide Hormones</i> , J. A. Parsons (ed.), University Park Press, Baltimore, pp. 1-7, 1976.					
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FORM PTO-449 (REV. 7-80)		MAR 8 4 2002 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE		ATTY. DOCKET NO. 690068.405C3		APPLICATION NO. 10/015,540	
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OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)							
MM	FA			Sakai et al., "Characterization of a major encephalitogenic T cell epitope in SJL/J mice with synthetic oligopeptides of myelin basic protein," <i>Journal of Neuroimmunology</i> 19: 21-32, 1988.			
	FB			Sakai et al., "Prevention of experimental encephalomyelitis with peptides that block interaction of T cells with major histocompatibility complex proteins," <i>Proc. Natl. Acad. Sci. USA</i> 86: 9470-9474, 1989.			
	FC			Santambrogio et al., "Altered peptide ligand modulation of experimental allergic encephalomyelitis: immune responses within the CNS," <i>Journal of Neuroimmunology</i> 81: 1-13, 1998.			
	FD			Seder and Paul, "Acquisition Of Lymphokine-Producing Phenotype By CD4 <sup>+</sup> T Cells," <i>Annu. Rev. Immunol.</i> 12: 635-673, 1994.			
	FE			Servis et al., "Two adjacent epitopes on a synthetic dodecapeptide induce lactate dehydrogenase B-specific helper and suppressor T cells," <i>Proc. R. Soc. Lond. B</i> 228: 461-470, 1986.			
	FF			Sette et al., "Analysis of lysozyme-specific immune responses by synthetic peptides. I. Characterization of antibody and T cell-mediated responses to the N-terminal peptide of hen egg-white lysozyme," <i>Eur. J. Immunol.</i> 16: 1-6, 1986.			
	FG			Sloan-Lancaster and Allen, "Significance of T-cell stimulation by altered peptide ligands in T cell biology," <i>Current Opinion in Immunology</i> 7: 103-109, 1995.			
	FH			Sloan-Lancaster et al., "Induction of T-cell anergy by altered T-cell-receptor ligand on live antigen-presenting cells," <i>Nature</i> 363: 156-159, 1993.			
	FI			Smilek et al., "A single amino acid change in a myelin basic protein peptide confers the capacity to prevent rather than induce experimental autoimmune encephalomyelitis," <i>Proc. Natl. Acad. Sci. USA</i> 88: 9633-9637, 1991.			
	FJ			Sriram et al., "Administration of Myelin Basic Protein-Coupled Spleen Cells Prevents Experimental Allergic Encephalitis," <i>Cellular Immunology</i> 75: 378-382, 1983.			
MM	FK			Steinman, "A few autoreactive cells in an autoimmune infiltrate control a vast population of nonspecific cells: A tale of smart bombs and the infantry," <i>Proc. Natl. Acad. Sci. USA</i> 93: 2253-2256, 1996.			
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FORM PTO-1492  
(REV. 7-80)U.S. DEPARTMENT OF COMMERCE  
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## OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

mm	GA	Steinman, "Autoimmune Disease. Misguided assaults on the self produce multiple sclerosis, juvenile diabetes and other chronic illnesses. Promising therapies are emerging," <i>Scientific American</i> 269(3): 106-114, September 1993.
	GB	Steinman, "Multiple Sclerosis: A Coordinated Immunological Attack against Myelin in the Central Nervous System," <i>Cell</i> 85: 299-302, 1996.
	GC	Steinman et al., "Major T-cell responses in multiple sclerosis," <i>Molecular Medicine Today</i> 1: 79-83, 1995.
	GD	Steinman et al., "Natural occurrence of thymocytes that react with myelin basic protein," <i>Neurology</i> 30(7): 755-759, 1980.
	GE	Steinman et al., "Regulation of autosensitisation to encephalitogenic myelin basic protein by macrophage-associated and soluble antigen," <i>Nature</i> 265: 173-175, 1977.
	GF	Steinman et al., "The Epigenetics Of Multiple Sclerosis: Clues to Etiology and a Rationale for Immune Therapy," <i>Annu. Rev. Neurosci.</i> 17: 247-265, 1994.
	GG	Su et al., "Synthetic Myelin Basic Protein Peptide Analogs Are Specific Inhibitors of Phospholipid/Calcium-Dependent Protein Kinase (Protein Kinase C)," <i>Biochemical and Biophysical Research Communications</i> 134(1): 78-84, 1986.
	GH	Talmadge et al., "Screening Models for Biological Response Modifiers," in <i>13<sup>th</sup> International Congress of Chemotherapy. Symposium, Biological Responses Modifiers; SY 64 part 203</i> , Vienna 8/28-9/2, 1993, pp. 203/19-203/34.
	GI	Teitelbaum et al., "Specific inhibition of the T-cell response to myelin basic protein by the synthetic copolymer Cop 1," <i>Proc. Natl. Acad. Sci. USA</i> 85: 9724-9728, 1988.
	GJ	Vogt et al., "Ligand Motifs of HLA-DRB5*0101 and DRB1*1501 Molecules Delineated from Self-Peptides," <i>Journal of Immunology</i> 153: 1665-1673, 1994.
	GK	Weigle, "Analysis of Autoimmunity through Experimental Models of Thyroiditis and Allergic Encephalomyelitis," <i>Advances In Immunology</i> 30: 159-273, 1980.
mm	GL	Wisniewski and Keith, "Chronic Relapsing Experimental Allergic Encephalomyelitis: An Experimental Model of Multiple Sclerosis," <i>Ann. Neurol.</i> 1: 144-148, 1977.

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Sheet 8 of 8

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mm	HA	Wraith et al., "Antigen Recognition in Autoimmune Encephalomyelitis and the Potential for Peptide-Mediated Immunotherapy," <i>Cell</i> 59: 247-255, 1989.
	HB	Wraith et al., "T Cell Recognition as the Target for Immune Intervention in Autoimmune Disease," <i>Cell</i> 57: 709-715, 1989.
	HC	Wucherpennig et al., "Structural Requirements for Binding of an Immunodominant Myelin Basic Protein Peptide to DR2 Isotypes and for Its Recognition by Human T Cell Clones," <i>J. Exp. Med.</i> 179: 279-290, 1994.
	HD	Zamvil and Steinman, "The T Lymphocyte in Experimental Allergic Encephalomyelitis," <i>Annu. Rev. Immunol.</i> 8: 579-621, 1990.
	HE	Zamvil et al., "Multiple Discrete Encephalitogenic Epitopes of the Autoantigen Myelin Basic Protein Include a Determinant for I-E Class II Restricted T Cells," <i>J. Exp. Med.</i> 168: 1181-1186, 1988.
	HF	Zamvil et al., "T-cell clones specific for myelin basic protein induce chronic relapsing paralysis and demyelination," <i>Nature</i> 317: 355-358, 1985.
	HG	Zamvil et al., "Encephalitogenic T Cell Clones Specific for Myelin Basic Protein," <i>J. Exp. Med.</i> 162: 2107-2124, 1985.
	HH	Zamvil et al., "T-cell epitope of the autoantigen myelin basic protein that induces encephalomyelitis," <i>Nature</i> 324: 258-260, 1986.
mt	HI	Zamvil et al., "T-cell Specificity for Class II(I-A) and the Encephalitogenic N-Terminal Epitope of the Autoantigen Myelin Basic Protein," <i>Journal of Immunology</i> 139(4): 1075-1079, 1987.
	HJ	
	HK	
	HL	
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